

1	CCACGCGGTC	CGCATAAATC	AGCAGCGGGC	CGGAGAACCC	CGCAATCTCT	GGCCCCACAA	AATACACCGA	CGATGCCCCG	TCCTACTTTAA	GGCTGAAC
	GGGTGGCGAG	CGGTATTTAG	TCGTGGCGCC	GCCTCTTGGG	CGGTATAGAG	CGCGGTGTT	TTATGTGGCT	GCTACGGGGT	AGATGAATT	CCGACTTTG
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	GGTCCCGGA	CTCTCTGATA	TTCTCGCAAG	GGATGGCGGT	ACCTTGTTC	CCCTGTCTTG	CGGGGCGCG	GRAGCCCGG	GGCTTTTCC	GTGCGGGTC
1										
201	GACCCAGGGA	GGGCGGGGA	GCCAGGCGTG	GGCTCGGGT	CCCCAGACC	CTGTGCTCG	TTGTGCGCG	GGTCTGCTG	TTGGTCTCAG	CTGAGTCTG
	CTGGTGCCT	CGGCGCCCT	CGGTCCGAC	CCGAGGCCA	GGGTTCCTG	GAACAGAGC	ACAGCGCG	CCAGGACGAC	AACAGAGTC	GACTCAGAC
22	Proarg1	uAlaArgGly	AlaArgProG	lyLeuArgVa	lProLysThr	LeuValLeuV	alValAlaAl	avalLeuLeu	LeuValSerA	laGluserAla
301	TCGTGATCAC	CAACAAGACC	TAGCTCCCA	GCAGAGAGCG	CCCCACAC	AAAGACGTC	CAGCCCTCA	GAGGGATTGT	GTCCACCTGG	ACACCATATC
	AGACTACTCG	GTGTCTCTG	ATCGAGGGT	CGTCTCTCG	TTTTCTCCAG	GTCCGGGAGT	GTCCCTAAC	CTCCCTAAC	CAGGTGGACC	TGTGGTATAG
55	LeuileThr	GlnGlnAspL	eAlaProG1	nGlnArgAla	AlaProGlnG	lnLysArgse	rSerProSer	GlUGlyLeuC	ysProProG1	yHisHisile
401	TCAGNAGACG	GTAGAGATTG	CATCTCTG	AAATATGGAC	AGGACTATAG	CACCTCACTG	AATGACCTCC	TTTTCTGCTT	GGCTGCCACC	AGGTGTGATT
	AGCTTCTG	CATCTCTAAC	GTAGAGGACG	TTATATACCTG	TCCTGATATC	GTGAGTGACC	TTACTGGAGG	AAAGACGAA	OGGACGTGG	TCCACACTAA
88	serGluaspG	lyArgaspCy	sileserCys	lystYrGlyG	lnAspTyse	rThrHisTrp	AsnaapLeuL	eupheCysLe	uArgCysThr	ArgCysaspSer
501	CAGGTGAAGT	GAGCTAAGT	CCCTGCACCA	CGACAGAAA	CACAGTGCTG	CAGTGGCAG	AAGGCACCTT	CGGGGAAGAA	GATTCTCTG	AGATGTGCCG
	GTCCACTTCA	CTCGATTCA	GGGACGTGGT	GCCTGCTTTT	GTGTACACA	GTACAGCTTC	TTCCGTGAA	GGCCCTTCTT	CTAAGACGAC	TCTACACGGC
122	GlyGluVa	lGluLeuser	ProCysThrT	hrThrArgAs	nThrValCys	GlnCysGluG	lUGlyThrPh	eArgGluGlu	AspserProG	lumeiCysArg
601	GAAGTCCCG	ACAGGTGTC	CCAGAGGGAT	GGTCAAGGTC	GGTATTTGTA	CACCCTGGAG	TGACATCGAA	TGTGTCCACA	AAGAATCAGG	CATCATCAT
	CTTACCGCG	TGTCCACAG	GGTCTCCCTA	CCAGTTCCAG	CCACTAACAT	GTGGGACCTC	ACTGTAGCTT	ACACAGGTGT	TTCTTAGTCC	GCTAGTATAT
155	LysCysArg	ThrGlyCysP	roArgGlyMe	valLysVal	GlyaspCyst	hrProTrpSe	rAspIleGlu	CysValHisL	ysGluserG1	yllelle
701	GGAGTCACAG	TTGCAGCCGT	AGTCTTGATT	GTGGCTGCTG	TGTGTTGCAA	GTCTTTTACTG	TGGAACAAAG	TCCTTCTCTTA	CCTGAAGGC	ATCTGCTCAG
	CCTCAGTGTC	AACTGCGCA	TCAGAACTAA	CACCGACACA	ACAAACGTT	CAGAAATGAC	ACCTTCTTTC	AGGAAGGAAT	GGACTTTCCG	TAGACGAGTC
188	GlyValThrV	alAlaAlaVa	lValLeuile	valAlaValP	heValCysly	sSerLeuLeu	TriplysLysV	alleuProTy	xLeuLysGly	lleCysSerGly
801	GTGGTGTGG	GGACCCGAC	CGTGTGGACA	GAACTCCACA	AGACCTGGG	GCTGAGGACA	ATGCTCTCAA	TGAGATCGTG	AGTATCTTTC	AGCCACCCA
	CACCAACACC	CTGGGACATC	GCACACCTGT	CTTCGAGTGT	TGCTGGACCC	CGACTCTCTG	TACAGGAGTT	ACTCTAGCAC	TCATAGNACG	TCGGGTGGGT
222	GlyGlyG1	ysAppProGlu	ArgValAspA	rgserSerG1	nArgProGly	AlaGluAspA	snValLeuAs	nclulleVal	SerlleLeuG	lnProThrGln
901	GTTCCTCTG	CAGGAAATGG	ARGTCCAGGA	GCCAGGCAG	CCAAACGCTG	TCAACATGTT	GTCCCCCGGG	GACTCAGAGC	ATCTGCTGGA	ACCGGCAGAA
	CCAGGGACTC	GTCTTTTACC	TTACAGGTCTC	CGGTCTCTC	GGTTGTCAC	AGTTGTACAA	CAGGGGGGCC	CTCAGTCTCG	TAGACGACCT	TGGCCGTCTT
255	ValProGlu	GlnGluMetG	luValGlnG1	uProAlaGlu	ProThrGlyV	AlasnMetLe	userProGly	GluserGluH	isLeuLeuG1	uProAlaGlu
1001	CTCAAAAGGT	CTCAGAGGG	GAGGTGCTG	GTTCCAGCAA	ATGAAGGTGA	TGCCACTGAG	ACTCTGAGAC	AGTCTTCTCA	TGACTTTTGA	GACTTGGTGC
	CGACTTTTCCA	GAGTCTCTC	CTCCAGCAC	CAAGGTCTGT	TACTTCCACT	AGGCTGACTC	TGAGACTCTG	TCACGAGCT	ACTGAACCT	CTGAACCCAG
288	AlaGluArgS	exGlnArgAr	gArgLeuLeu	valProAlaA	snGluGlyAs	pProThrGlu	ThrLeuArgG	lnCysPheAs	pasPpheAla	AspleuValPro

1101 CCTTTGACTC CTGGAGCCG CTCATGAGGA AGTTGGGCGCT CATGCAAT GAGATAAGG TGGCTAAGG TGAGCAGCG GGCCACAGGG ACACCTTGT
 322 GAAACCTAG GACCTCGG GAGTACTCCT TCAACCCGGA GTACCTCTTA CTCTATTTC ACGATTTCG ACTGCTGCG CCGCTGCCC TGTGGAACAT
 pheAsp8 rTTPGluPro LeuMetArgL ysLeuGlyLe uMetAspAn GluileLysV alalalysAl aGluAlaLa GlyHisArga spThrLeuTyr
 1201 CAGCATGCTG ATAAAGTGG TCAACAAAC CGGCGAGAT GCTCTCTCC ACACCTGCT GGATGCTTG GAGAGAGACT TGCCAAAGCAG
 355 ThrMetLeu IleLysTrp alaAsnLysTh rGlyArgAsp AlaSerValH leThrLeuLe uAspAlaLeu GluThrLeuG lyGluArgLe uAlaLysGln
 1301 AAGATTGAGG ACACCTGCT GAGCTCTGA AAGTTATGT ATCTAGAGG TAATGAGAC TCTGCCWGT CCTAAGTGTG ATCTCTTCA GGAAGTGAGA
 388 LysIleGluA spHisLeuLe uSerSerGly LysPheMetT yrLeuGluG lYAsnAlaAsp SerAlaXqqS erDG*
 1401 CCTTCCCTGG TTACCTTTT TTCTGGA AAA AGCCCACTG GACTCCAGTC AGTAGGAAAG TGCCACAATT GTACATGAC CGCTACTGGA AGAACTCTC
 GGAAGGGACC AATGGGAAA AACACCTTTT TGGGCTGAC CTGAGGTCAG TCATCTTTTC ACGGTGTAA CAGTGTACTG GCCATGACCT TCTTTGAGAG
 1501 CCATCCAAAC TCACCCAGTG GATGGAACAT CCTGTAACTT TTCACTGCAC TTGGCATTTAT TTTTATAAGC TGAATGTGAT AATAAGGACA CTATGGAAT
 GGTAGGTGT AGTGGGTGAC CTACCTTGTGTA GGACATTGAA AAGTGACCTG AACCGTAATA AAAATATTG ACTTACACTA TTATTCTCTG GATACCTTTA
 1601 GTCTGGATCA TTCCGTTGT GCGTACTTTG AGATTGGTT TGGGATGTCA TTGTTTTAC ACCACTTTTT ACCACTTTTT TATCCTAATG TAAATGCTTT ATTATTTAT
 CAGACCTAGT AAGGCNAACA CGCATGAAAC TCTAAACCAA ACCCTACAGT AACRAAAGTC TCGTGA AAA ATAGGATTAC ATTTACGAAA TAAATPAAATA
 1701 TTGGGGTACA TTGTAGATC CATCTACAA AAAAAAAA AAAAAAAG GCGGCGCGG ACTCTAGAGT CGACCTGCAG AAGCTTGGCC GCCATGGCC
 AACCCGATGT AACATTCTAG GTAGATGTTT TTTTTTTTTT TTTTTTTTTT CCGCGGCGG TCAGATCTCA CCTGGACGTC TTGGAACCGG CCGTACCGG

Fig. 1 (cont.)

665760" OF 296600

ig. 2 A

1 MEORGONAPASGARKRHGPGPREARGARGLRVPKTLVLVAALLLVSAESALITQOD
61 LAPQORAPQOKRSSPSEGLCPGPHRISEGDHCISCKYGQDYSTHWNLLFCRCRTRCD
121 SCEVELSPCTTIBNTVQCCEGTFREEDSPENCRKRTGCPRGMVKVGDCTPWSDI ECVH
181 KESGIIIGVTVAARVLIIVAVFCKSLMKKVLPLYKGCISGGGDPERVDRSSORPGAED
241 NVLNEIVSILQPTQVPEQEMEVOEPAEPTGVNMLSPGESEHLLERAEAEHSQRRLLVPA
301 NEGDPTEITLROCFDDFADILVFFDSWEPLMRKLGMDNEIKVAKAEFAAGHRDITLYTMLIKW
361 VNKTRGDASVHTLLDALETGLERLAKQKIEDHLLSSGKFMYLEGNADSALS

Apo2
DR4
Apo3/DR3
TNFR1
Fas/Apo1

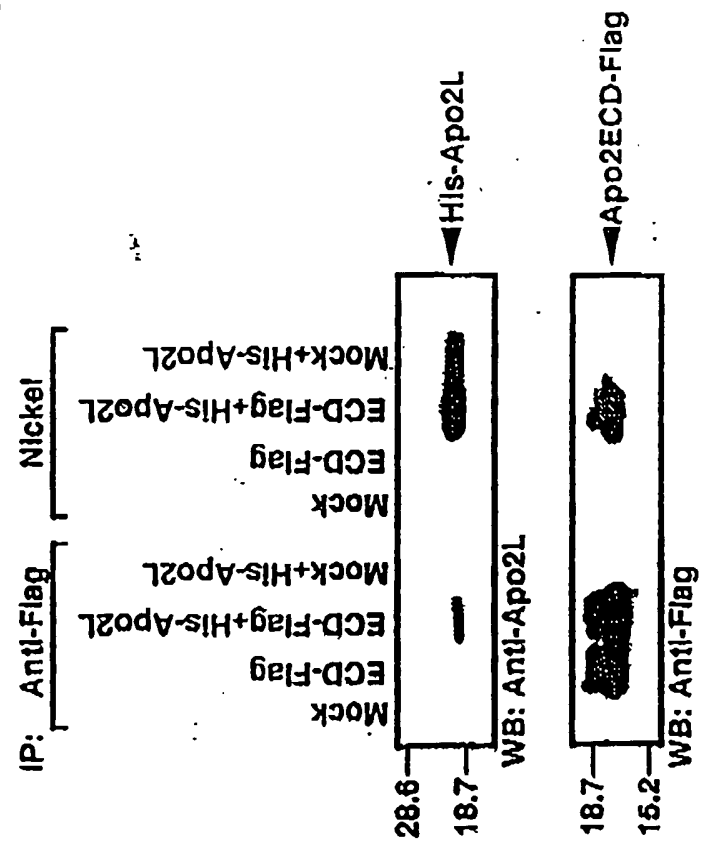
ig. 2 B

1 MEORGONAPASGARKRHGPGPREARGARGLRVPKTLVLVAALLLVSAESALITQOD
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121 SCEVELSPCTTIBNTVQCCEGTFREEDSPENCRKRTGCPRGMVKVGDCTPWSDI ECVH
181 KESGIIIGVTVAARVLIIVAVFCKSLMKKVLPLYKGCISGGGDPERVDRSSORPGAED
241 NVLNEIVSILQPTQVPEQEMEVOEPAEPTGVNMLSPGESEHLLERAEAEHSQRRLLVPA
301 NEGDPTEITLROCFDDFADILVFFDSWEPLMRKLGMDNEIKVAKAEFAAGHRDITLYTMLIKW
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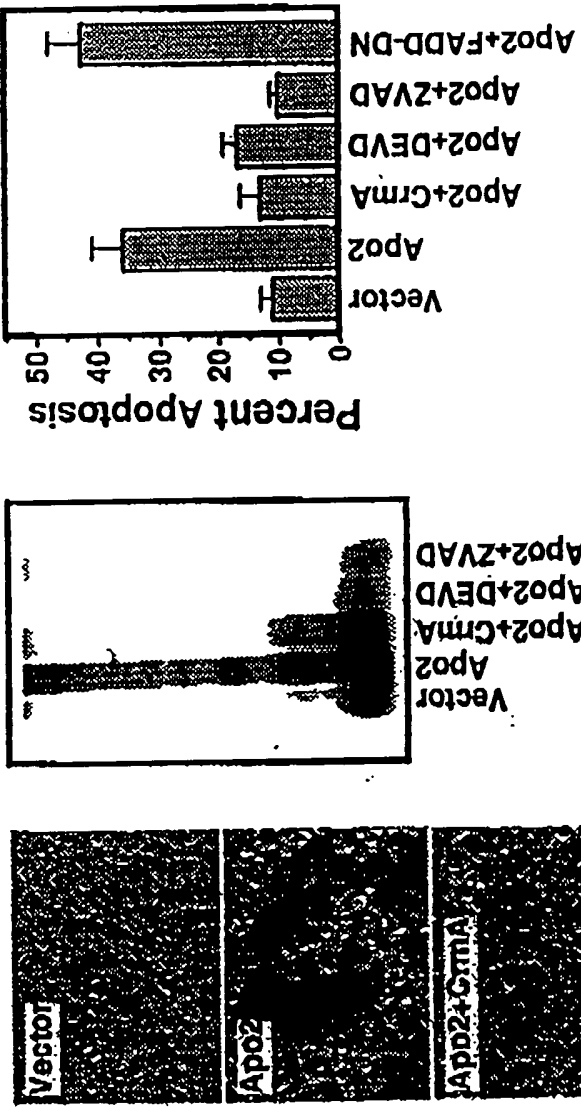
Apo2
DR4
Apo3/DR3
TNFR1
Fas/Apo1

1 MEORGONAPASGARKRHGPGPREARGARGLRVPKTLVLVAALLLVSAESALITQOD
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121 SCEVELSPCTTIBNTVQCCEGTFREEDSPENCRKRTGCPRGMVKVGDCTPWSDI ECVH
181 KESGIIIGVTVAARVLIIVAVFCKSLMKKVLPLYKGCISGGGDPERVDRSSORPGAED
241 NVLNEIVSILQPTQVPEQEMEVOEPAEPTGVNMLSPGESEHLLERAEAEHSQRRLLVPA
301 NEGDPTEITLROCFDDFADILVFFDSWEPLMRKLGMDNEIKVAKAEFAAGHRDITLYTMLIKW
361 VNKTRGDASVHTLLDALETGLERLAKQKIEDHLLSSGKFMYLEGNADSALS

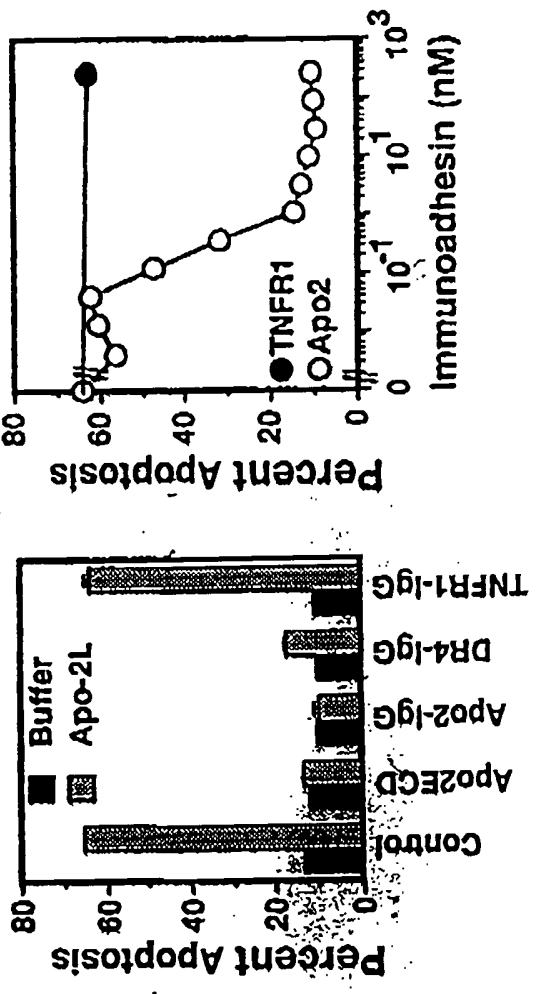
Fig. 3



4A 4B 4C

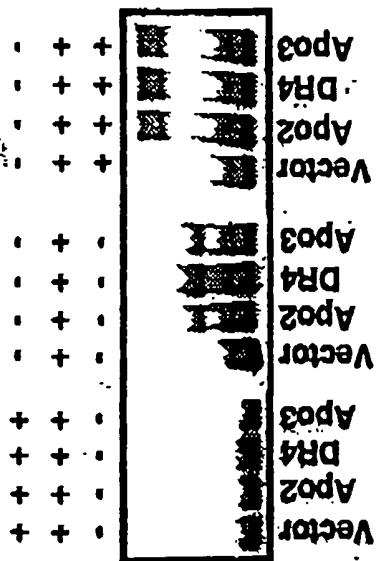


4D 4E

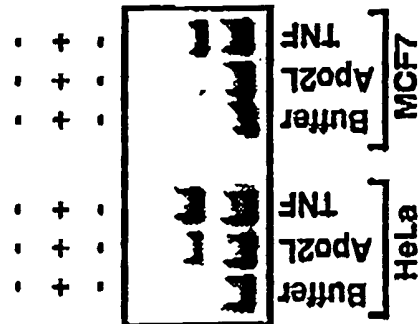


5A

Unlabelled probe
Labelled probe
Anti-p65



5B



5C

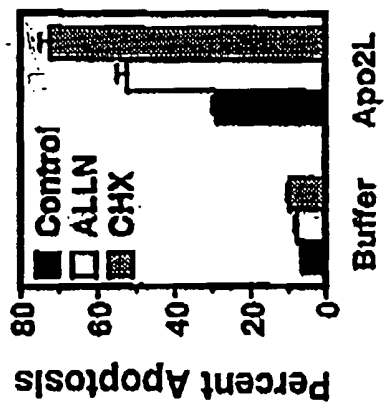
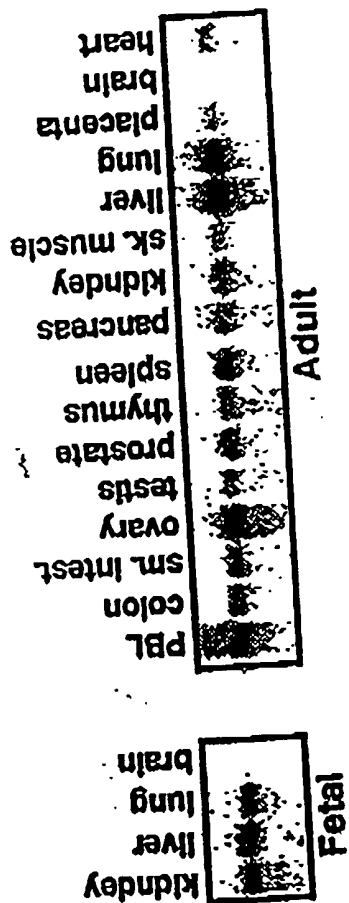


Fig. 5

FIG. 6



655T60" 07/96E60

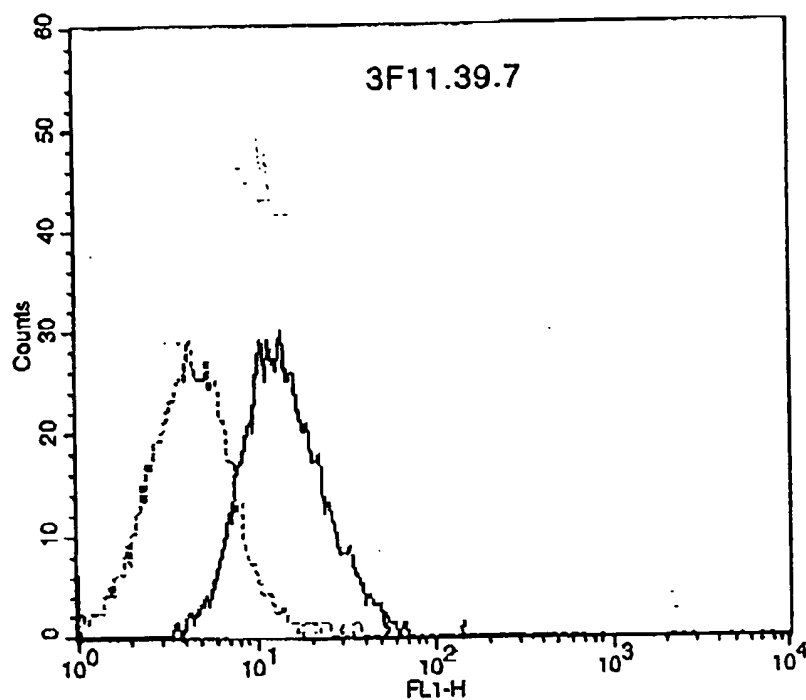


Fig. 7

005160" 01296260

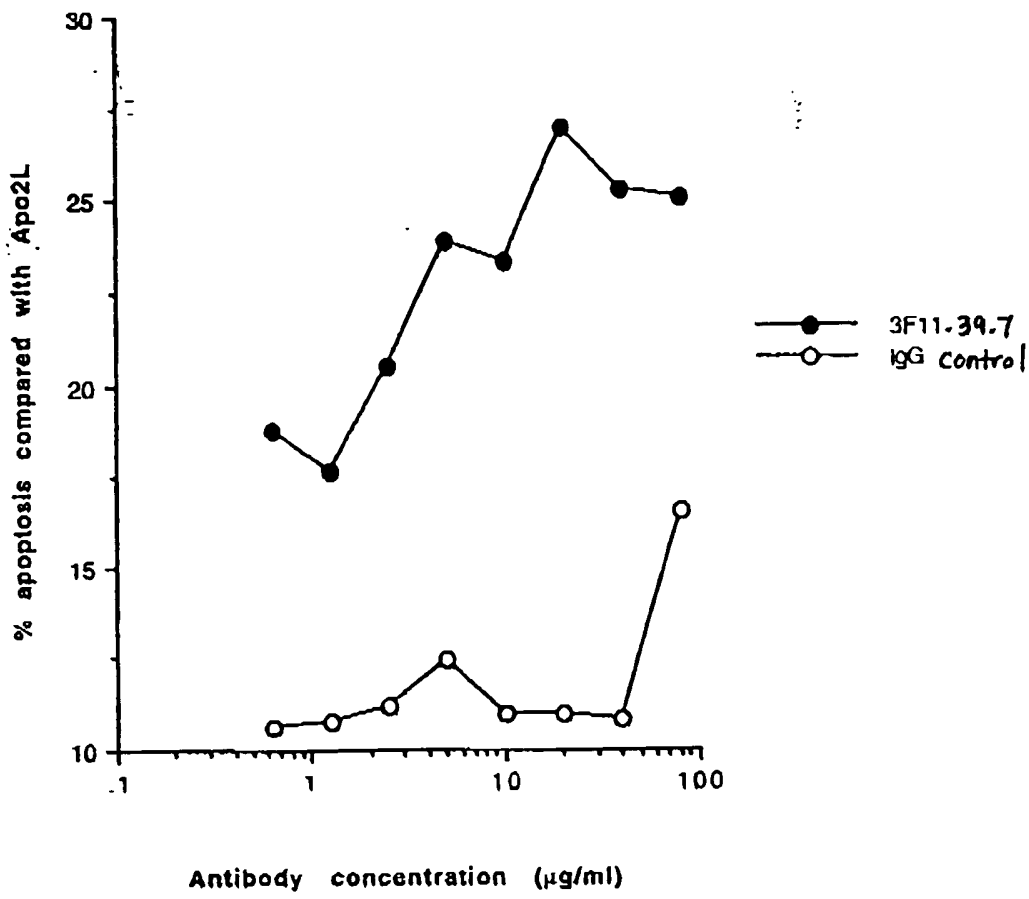


Fig. 8

09396710-09559

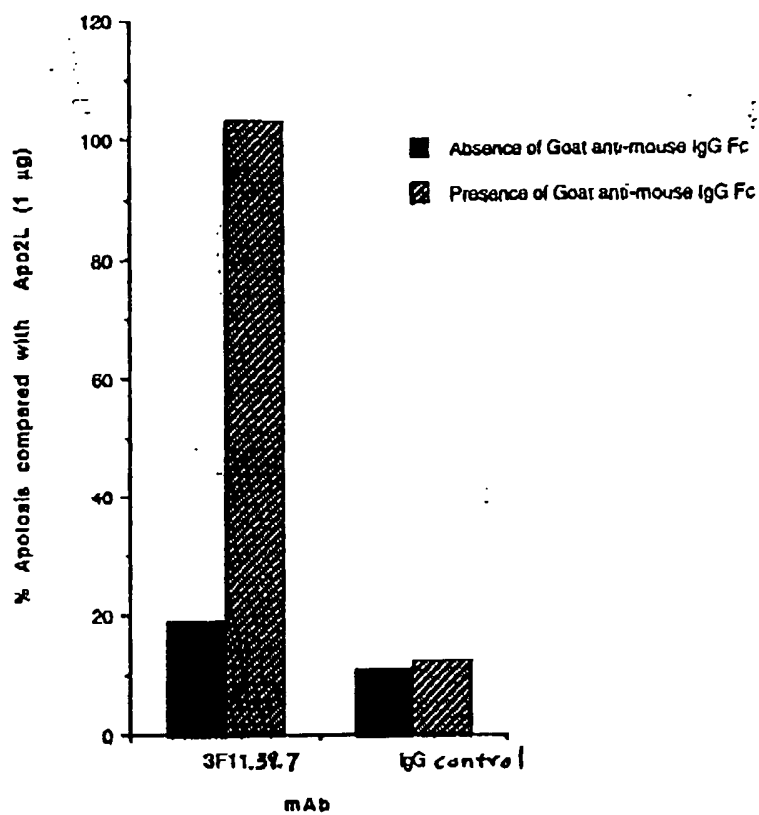


Fig. 9

09296710.091599

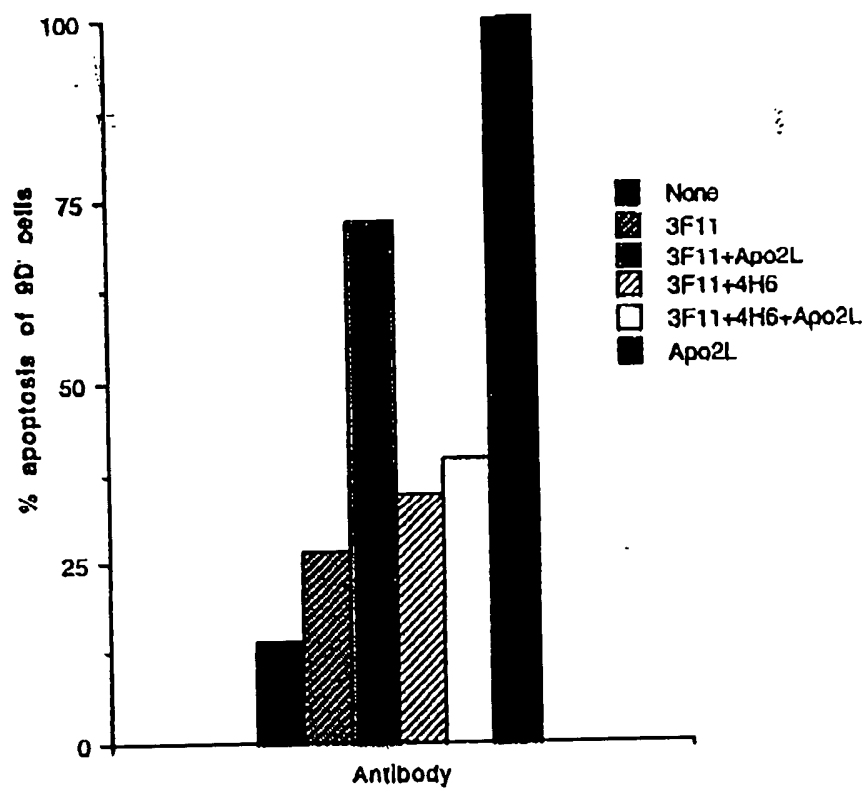


Fig : 10

665160" OF 296660

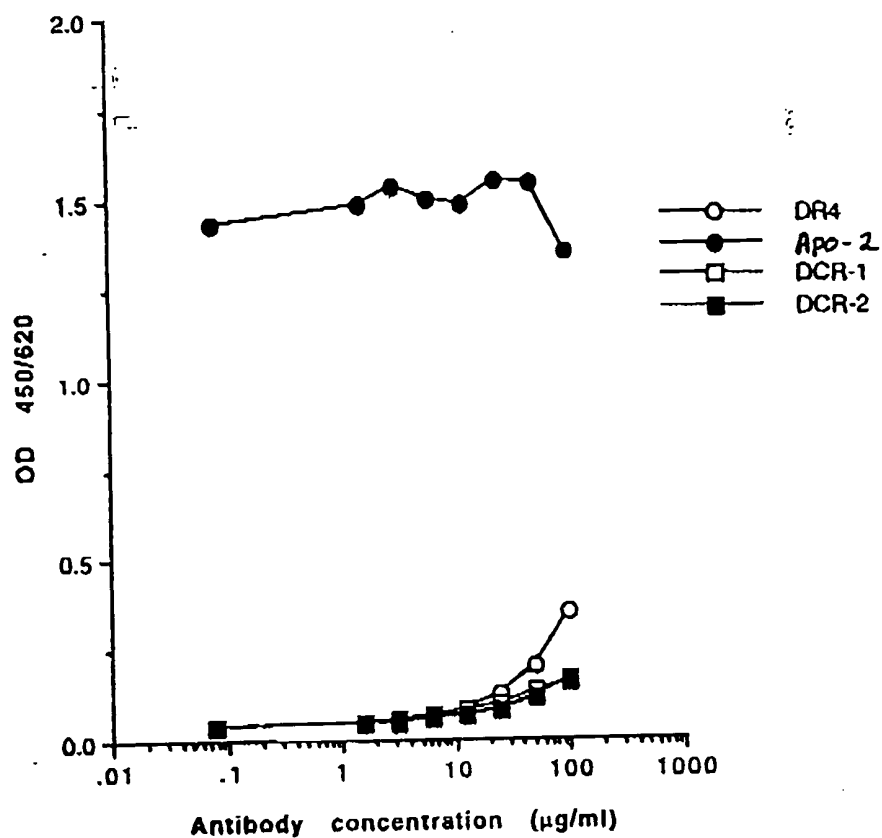


Fig. 11

Isotype	Kd-1 (pM)	Bloc	Apop w/L	Cross reactivity			IHC
				DR4	Apo-2	DCR1 DCR2	
2B3.7.1	20.	-	+	+	+++	-	+
3F11.39.7		+	+	-	+++	-	
4B9.23.6		-	+/-	-	+++	-	
5C7.9.1		-	+	+	+++	-	
3H1.18.10	4	-	-	+	+++	+/-	+
3H3.14.5		+	+	+	+++	+	+
3D5.1.10		-	+	++	+++	-	
3C9.8.6		+	+	-	+++	-	
4H10.14.10		+/-	?	+/-	+++	+/-	

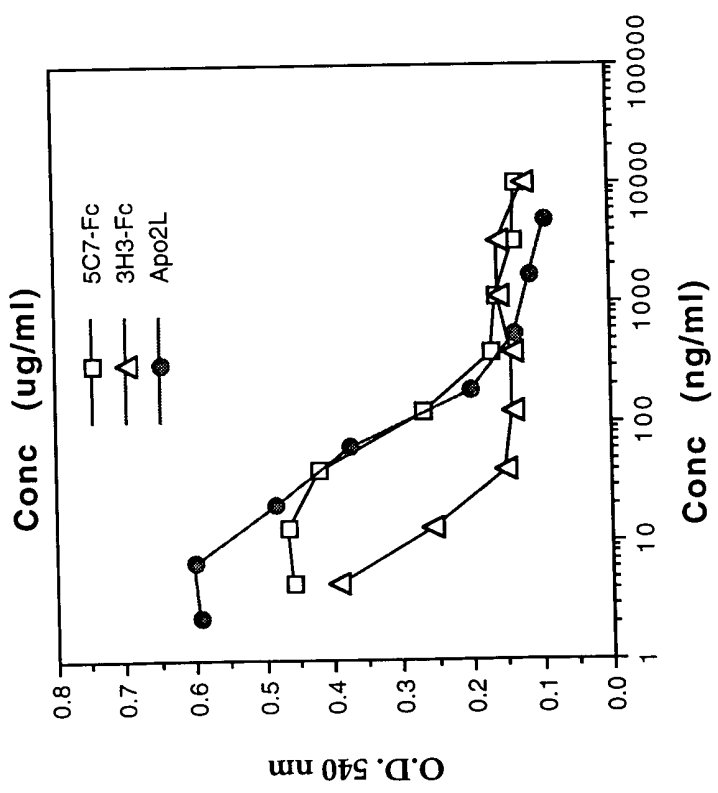
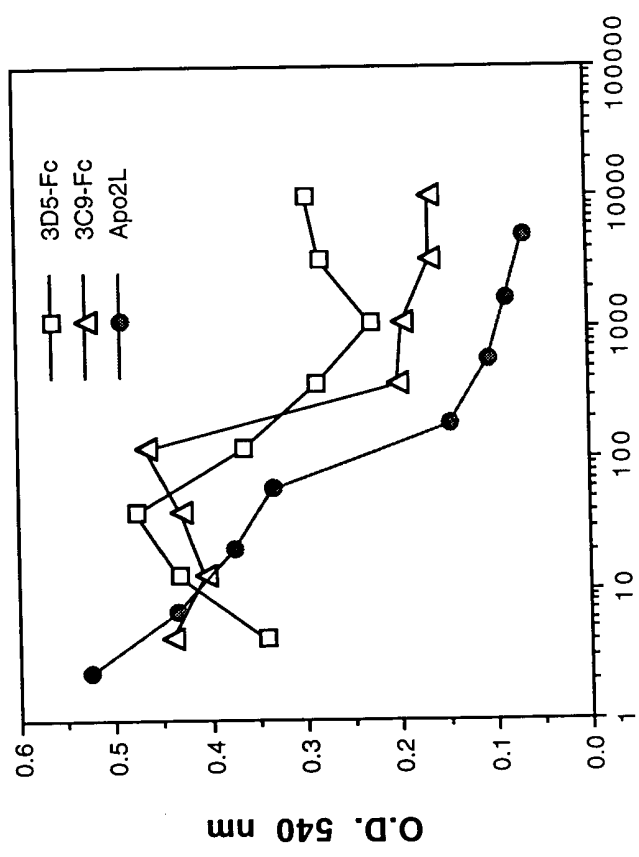
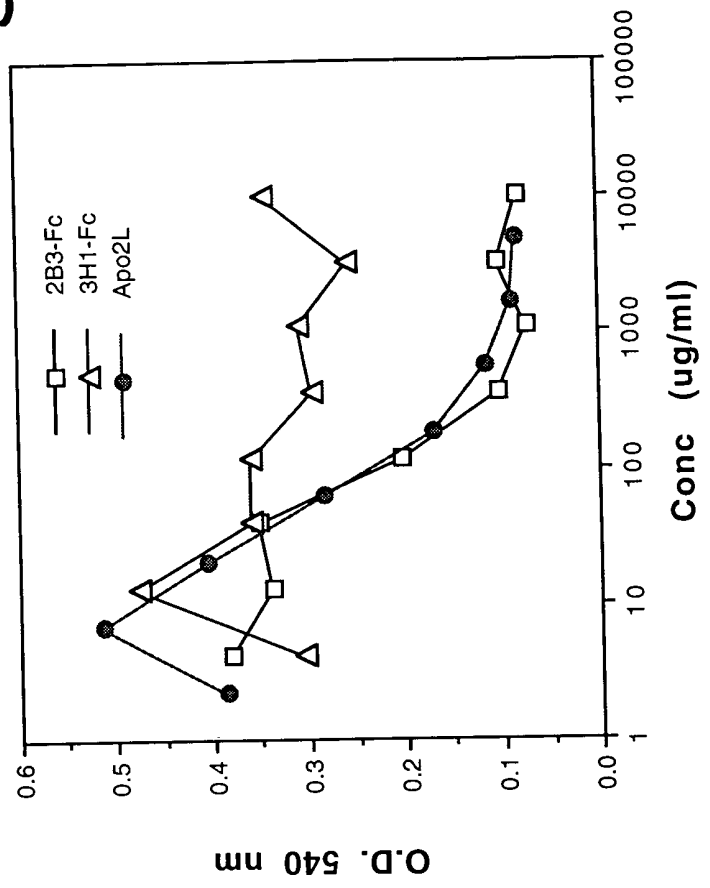
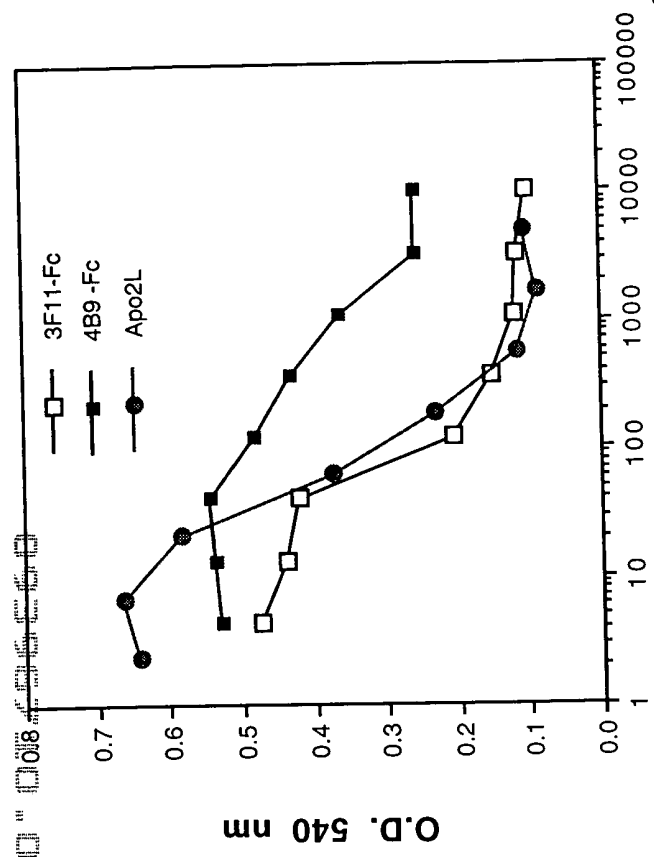
Fig. 12

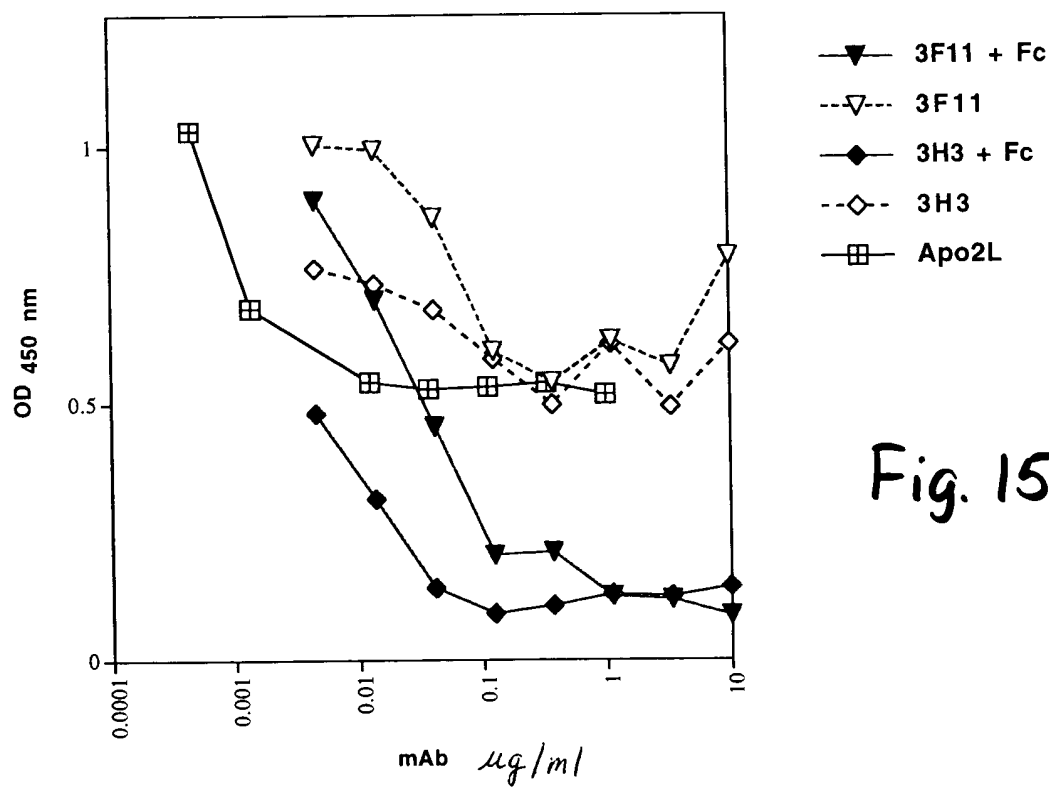
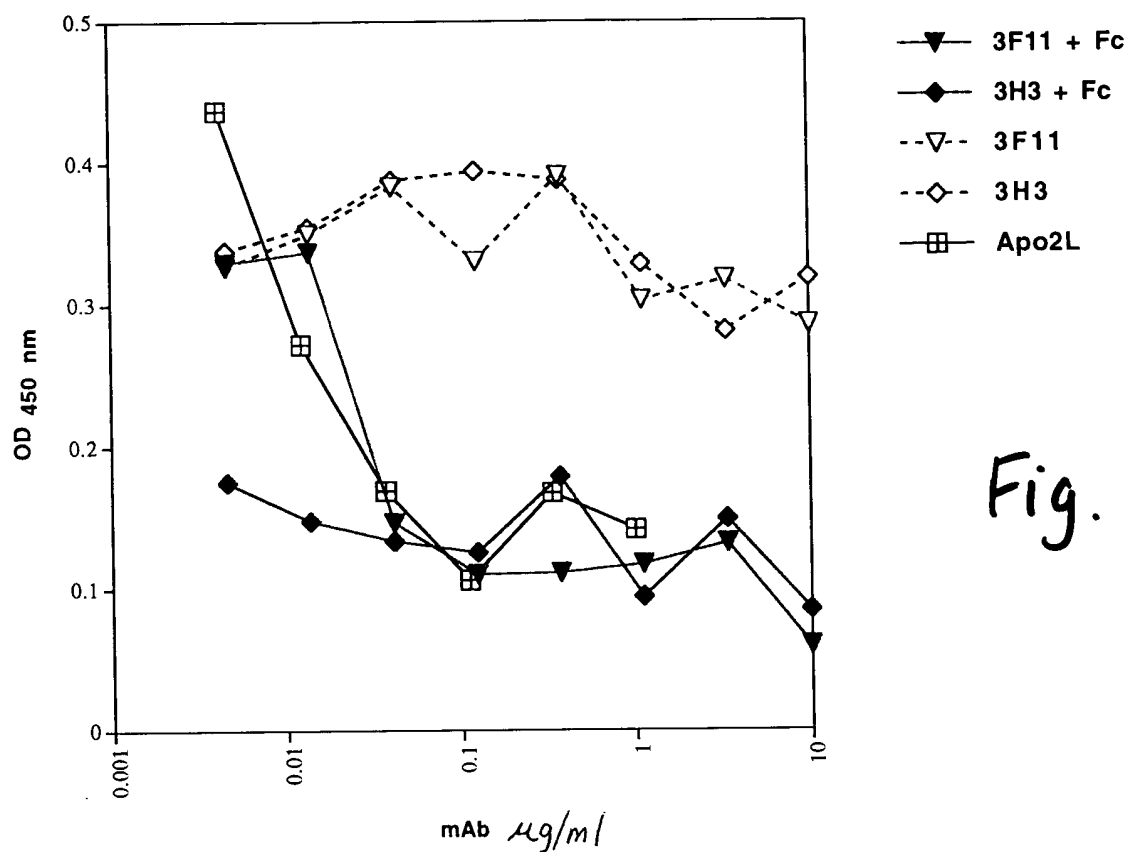
		Affinity (pM)
DR4-IgG	to Apo2L	82
Apo-2 -IgG	to Apo2L	1
mAb 3F11	to Apo-2 IgG	20
mAb 3H3	to "	3

Fig. 13

09396710 " 091599

Fig. 14





665160" DT 96660

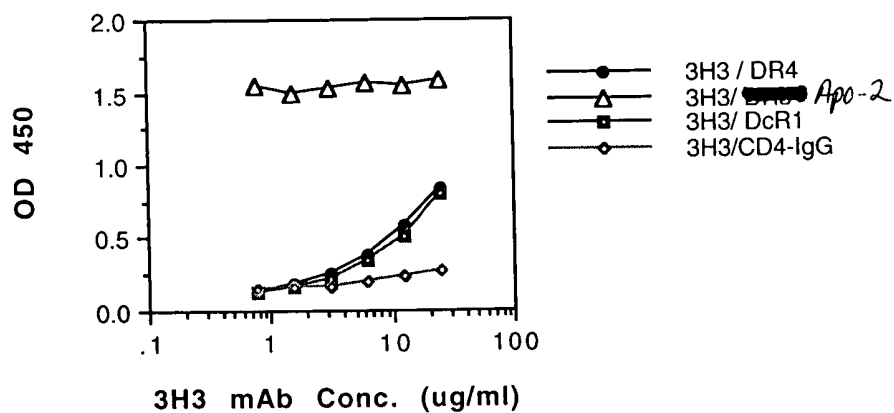
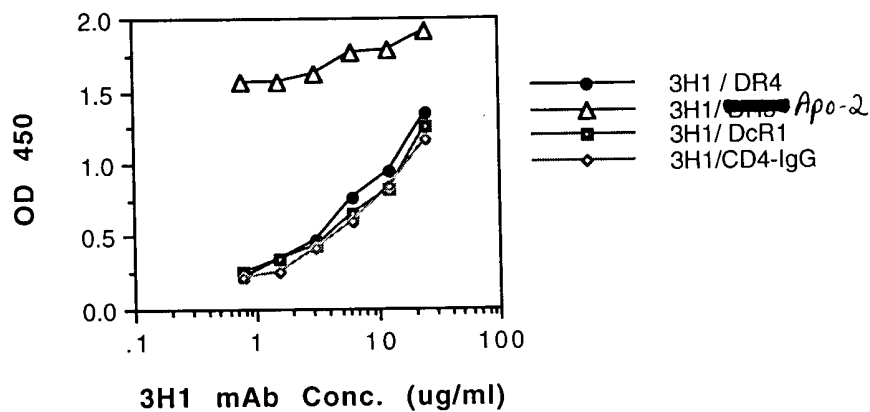
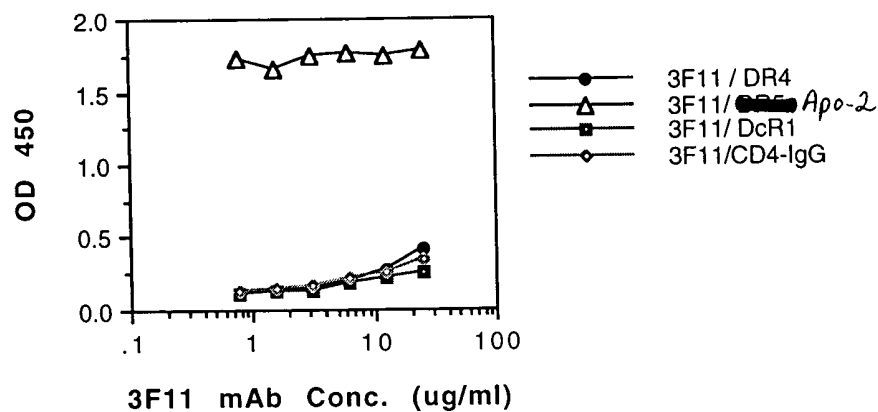
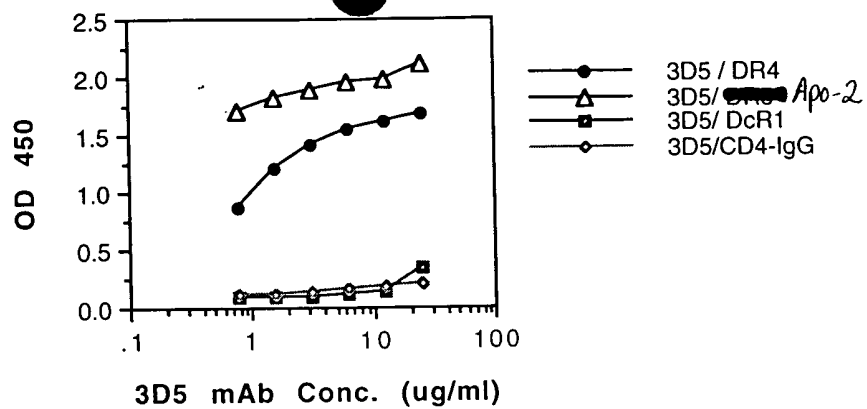


Fig. 16

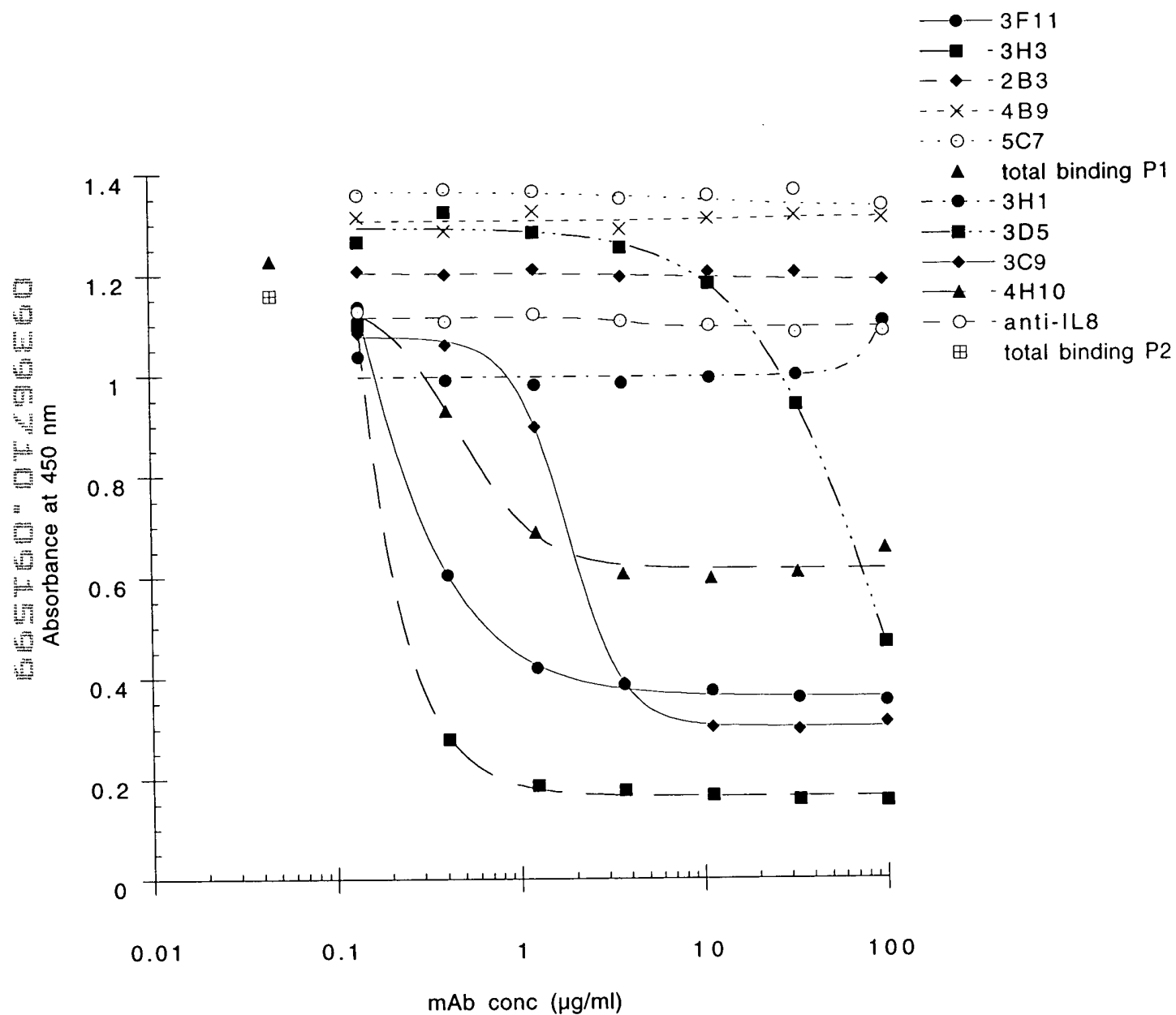


Fig. 17

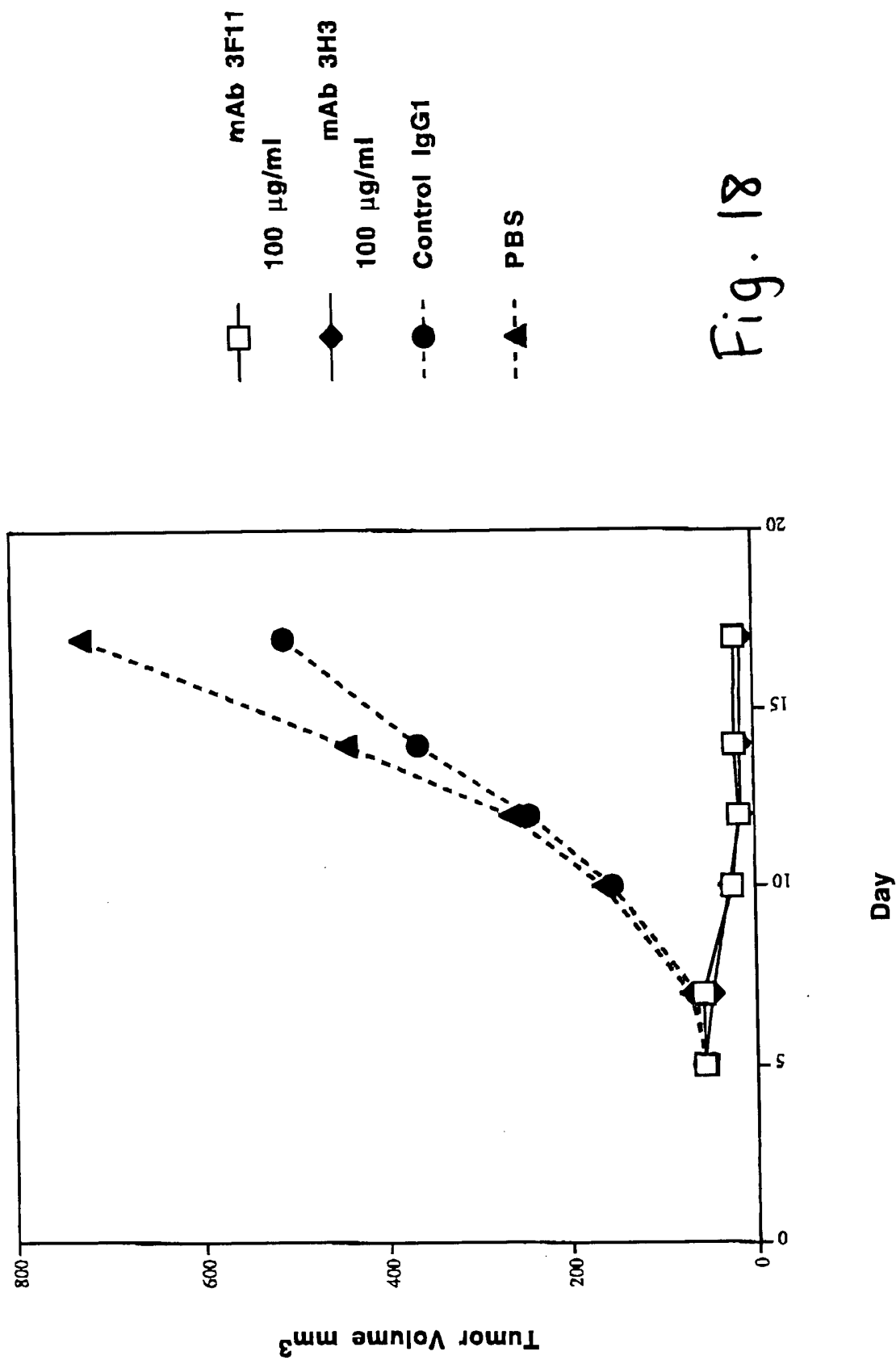


Fig. 18

Data Collection

	In-house	SSRL
Resolution (Å)	30-3.5 (3.62-3.50) ^a	30-2.4 (2.49-2.40) ^a
R _{sym} ^b	0.185 (0.398) ^a	0.056 (0.396) ^a
Number of observations	51,527	152,986
Unique reflections	12,459	38,908
Completeness (%)	99.8 (99.9) ^a	99.8 (99.7) ^a

Refinement

Resolution (Å)	30-2.4
Number of reflections	38,850
Final R ^c , R _{free} (F>0)	0.222, 0.267
Number of residues	781
Number of solvent molecules	286
Number of non-H atoms	6577
Average B factor (Å ²)	47.6
Rmsd bonds (Å)	0.013
Rmsd angles (°)	1.7
Rmsd B (bonded atoms) (Å ²)	2.4

^a Numbers in parentheses refer to the highest resolution shell.

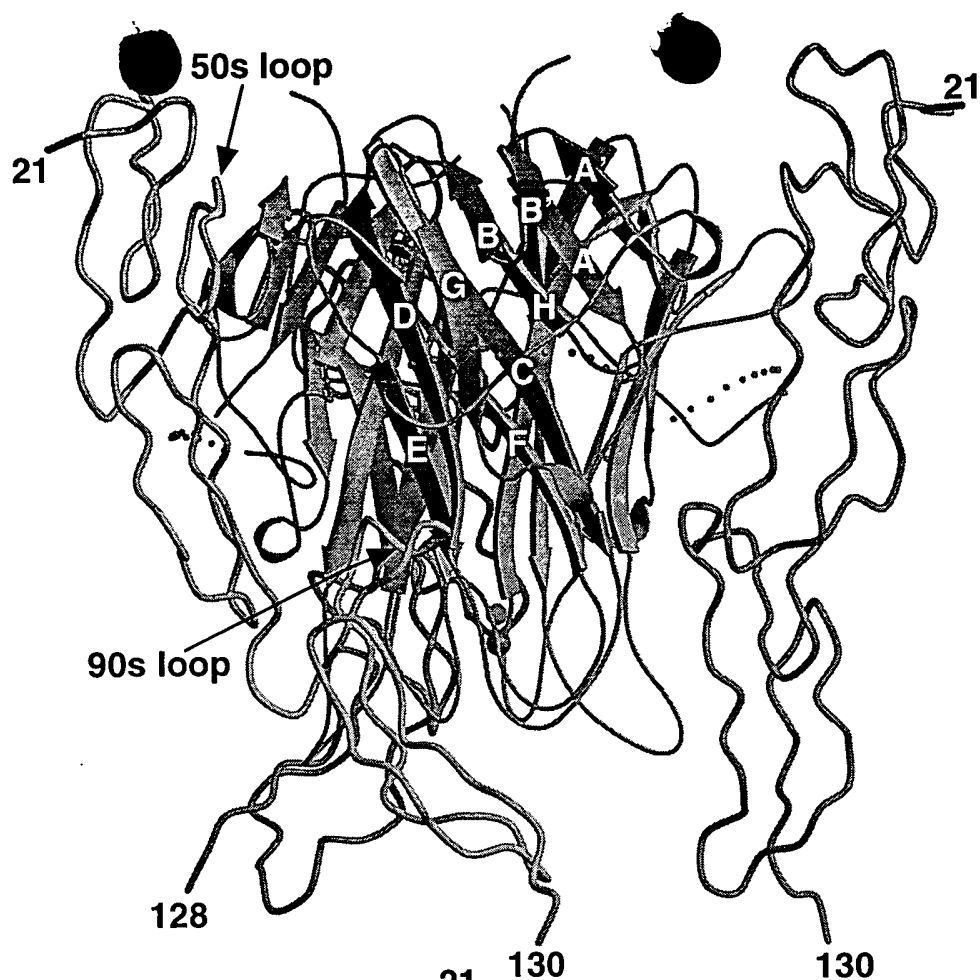
^b $R_{sym} = \sum ||I - \langle I \rangle| / \sum I$. $\langle I \rangle$ is the average intensity of symmetry related observations of a unique reflection.

^c $R = \sum |F_o - F_c| / \sum F_o$. R_{free} is calculated as R, but for 10% of the reflections excluded from all refinement.

Fig. 19

09396710.091569

A



B

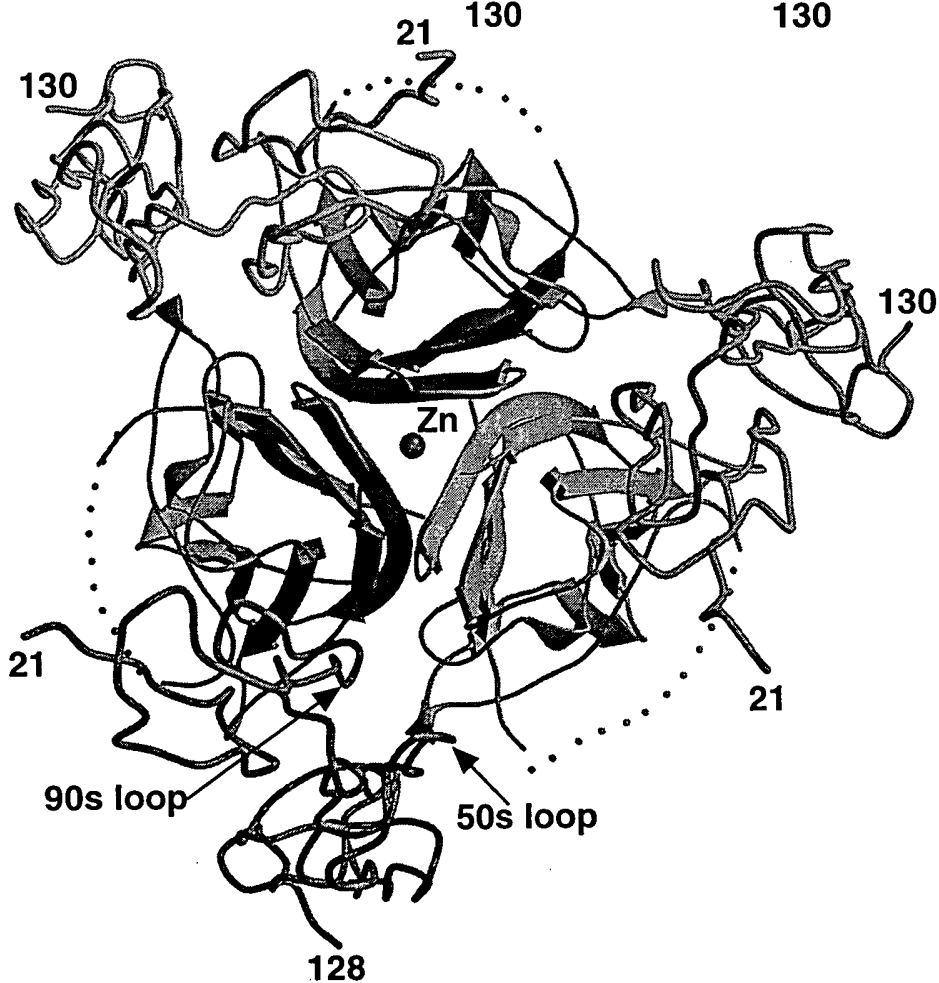


Figure 20

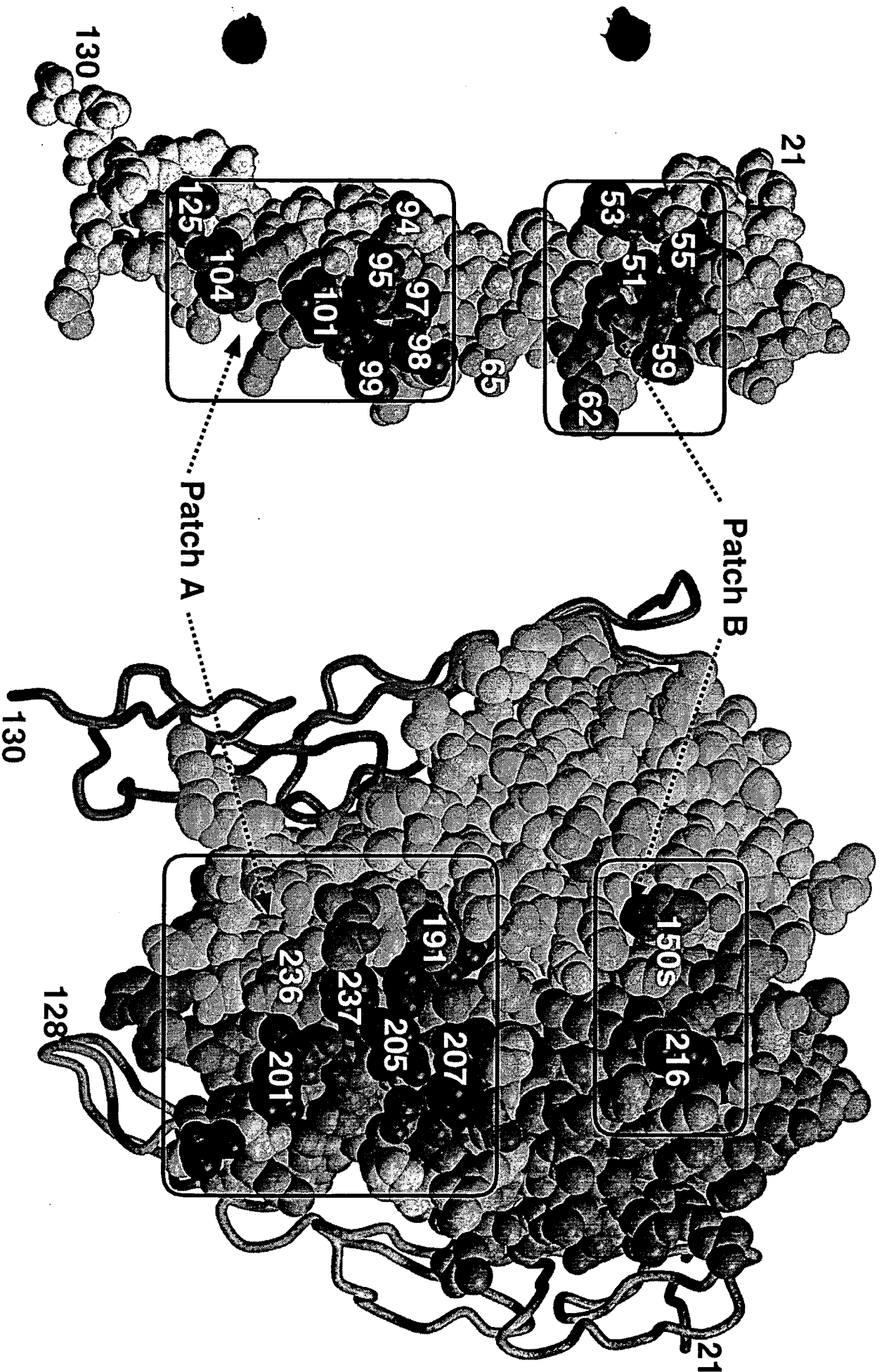


Figure 21